

### SEQUENCE LISTING

## (1) GENERAL INFORMATION:

(i) APPLICANT: Le, Junming Vilcek, Jan Daddona, Peter E. Ghrayeb, John Knight, David M.

(ii) TITLE OF INVENTION: METHODS OF TREATING TNFα-MEDIATED DISEASE USING CHIMERIC ANTI-TNF ANTIBODIES (As Amended)

## (iii) NUMBER OF SEQUENCES: 19

- (iv) CORRESPONDENCE ADDRESS:
  - (A) ADDRESSEE: Hamilton, Brook, Smith & Reynolds, P.C.
  - (B) STREET: Two Militia Drive

Siegel, Scott A.

- (C) CITY: Lexington
- (D) STATE: Massachusetts
- (E) COUNTRY: USA (F) ZIP: 02173
- (v) COMPUTER READABLE FORM: (A) MEDIUM TYPE: Floppy disk
  - (B) COMPUTER: IBM PC compatible
  - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
    - (D) SOFTWARE: PatentIn Release #1.0, Version #1.25

REUMANO

JUL 1 6 1998

**GROUP 1800** 

- (vi) CURRENT APPLICATION DATA:
  - (A) APPLICATION NUMBER: 08/192,861
  - (B) FILING DATE: 04-FEB-1994
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: US 08/013,413
  - (B) FILING DATE: 02-FEB-1993
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: US 08/010,406
    (B) FILING DATE: 29-JAN-1993
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: US 07/943,852
  - (B) FILING DATE: 11-SEP-1992
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: US 07/853,606
  - (B) FILING DATE: 18-MAR-1992
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: US 07/670,827
  - (B) FILING DATE: 18-MAR-1991

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- (viii) ATTORNEY/AGENT INFORMATION:
  - (A) NAME: Brook, David E.
  - (B) REGISTRATION NUMBER: 22,592
  - (C) REFERENCE/DOCKET NUMBER: NYU93-01M2
  - (ix) TELECOMMUNICATION INFORMATION:
    - (A) TELEPHONE: (781) 861-6240
    - (B) TELEFAX: (781) 861-9540
- (2) INFORMATION FOR SEQ ID NO:1:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 157 amino acids
      - (B) TYPE: amino acid
      - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

  - Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg Arg 20 25 30
  - Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln Leu
    35 40 45
  - Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu Phe 50 55 60
  - Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr Ile 65 70 75 80
  - Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser Ala 85 90 95
  - Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala Lys 100 \$105\$
  - Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu Lys 115 120 125
  - Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp Phe 130 135 140
  - Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155
- (2) INFORMATION FOR SEO ID NO:2:





	(i)	() ()	QUENCA) LE	engti (PE : [RANI	H: 32 nucl	21 ba leic ESS:	ase p acid	pairs 1	3					
	(ii)	MOI	LECUI	LE T	YPE:	CDN	Ą							
	(ix)	(1	ATURI A) NJ 3) LO	AME/I			321							
	(xi	) SE(	QUEN	CE DI	ESCR	PTI	ON: S	SEQ :	ID N	0:2:				
			CTG Leu										GGA Gly	48
			AGT Ser 20											96
			TAT Tyr										ATA Ile	144
			TCT Ser										GGC Gly	192
			GGG Gly											240
			GCA Ala											288
			TCG Ser 100											321
(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO : 3	:						
	(i) SEQUENCE CHARACTERISTICS: (a) LENGTH: 107 amino acids													

(B) TYPE: amino acid (D) TOPOLOGY: linear

(ii)	MOLECULE	TYPE:	protein
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ı	vi	) SECUENC	E DESCRIPTION:	SEO	TD	NO · 3 ·

Asp Ile Leu Leu Thr Gln Ser Pro Ala Ile Leu Ser Val Ser Pro Gly 1 5 10 15

Glu Arg Val Ser Phe Ser Cys Arg Ala Ser Gln Phe Val Gly Ser Ser 20 25 30

Ile His Trp Tyr Gln Gln Arg Thr Asn Gly Ser Pro Arg Leu Leu Ile  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Lys Tyr Ala Ser Glu Ser Met Ser Gly Ile Pro Ser Arg Phe Ser Gly 50 60

Ser Gly Ser Gly Thr Asp Phe Thr Leu Ser Ile Asn Thr Val Glu Ser 65 70 75 80

Glu Asp Ile Ala Asp Tyr Tyr Cys Gln Gln Ser His Ser Trp Pro Phe  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Thr Phe Gly Ser Gly Thr Asn Leu Glu Val Lys

# (2) INFORMATION FOR SEQ ID NO:4:

#### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 357 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
  (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: cDNA

# (ix) FEATURE:

- (A) NAME/KEY: CDS
- (B) LOCATION: 1..357

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

GAA GTG AAG CTT GAG GAG TCT GGA GGA GGC TTG GTG CAA CCT GGA GGA 48 Glu Val Lys Leu Slu Glu Ser Gly Gly Leu Val Gln Pro Gly Gly 1 10 15

TCC ATG AAA CTC TCC TGT GTT GCC TCT GGA TTC ATT TTC AGT AAC CAC 96 Ser Met Lys Leu Ser Cys Val Ala Ser Gly Phe Ile Phe Ser Asn His 20 25 30

TGG ATG AAC TGG GTC CGC CAG TCT CCA GAG AAG GGG CTT GAG TGG GTT 144 Trp Met Asn Trp Val Arg Gln Ser Pro Glu Lys Gly Leu Glu Trp Val  $\frac{1}{3}$ 5 40 45

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															GAG	192
Ala	Glu	Ile	Arg	Ser	Lys	ser	Ile	Asn	Ser	Ala	Thr	His	Tyr	Ala	Glu	
	50					55					60					
TCT	GTG	AAA	GGG	AGG	TTC	ACC	ATC	TCA	AGA	GAT	GAT	TCC	AAA	AGT	GCT	24
Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asp	Ser	Lys	Ser	Ala	
65			-	_	70					75					80	
GTC	TAC	CTG	CAA	ATG	ACC	GAC	TTA	AGA	ACT	GAA	GAC	ACT	GGC	GTT	TAT	28
Val	Tyr	Leu	Gln	Met	Thr	Asp	Leu	Arg	Thr	Glu	Asp	Thr	Gly	Val	Tyr	
	•			85					90					95		
TAC	TGT	TCC	AGG	AAT	TAC	TAC	GGT	AGT	ACC	TAC	GAC	TAC	TGG	GGC	CAA	33
Tyr	Cys	Ser	Arg	Asn	Tyr	Tyr	Gly	Ser	Thr	Tyr	Asp	Tyr	Trp	Gly	Gln	
-	-		100					105					110			

#### (2) INFORMATION FOR SEO ID NO:5:

GGC ACC ACT CTC ACA GTC TCC

Gly Thr Thr Leu Thr Val Ser

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 119 amino acids
  - (B) TYPE: amino acid
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:
- Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly 1 5 10 15
- Ser Met Lys Leu Ser Cys Val Ala Ser Gly Phe Ile Phe Ser Asn His 20 25 30
- Trp Met Asn Trp Val Arg Gln Ser Pro Glu Lys Gly Leu Glu Trp Val 35 40 45
- Ala Glu Ile Arg Ser Lys Ser Ile Asn Ser Ala Thr His Tyr Ala Glu 50  $\,$
- Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Ser Ala 65 70 75 80
- Val Tyr Leu Gln Met Thr Asp Leu Arg Thr Glu Asp Thr Gly Val Tyr 85 90 95
- Tyr Cys Ser Arg Asn Tyr Tyr Gly Ser Thr Tyr Asp Tyr Trp Gly Gln  $100 \hspace{1cm} 105 \hspace{1cm} 110 \hspace{1cm}$

Gly Thr Thr Leu Thr Val Ser 115

- (2) INFORMATION FOR SEQ ID NO:6:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 8 amino acids
    - (B) TYPE: amino acid
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: protein
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Gly Thr Leu Val Thr Val Ser Ser

- (2) INFORMATION FOR SEQ ID NO:7:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 7 amino acids
    - (B) TYPE: amino acid
    - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

Gly Thr Lys Leu Glu Ile Lys

- (2) INFORMATION FOR SEQ ID NO:8:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 20 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: cDNA
    - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

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CCTGGATACCTGTGAAAAGA

(2) INFORMATION FOR SEQ ID NO:9:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 27 base pairs
  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single

  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA



(xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:	
CCTGGTACCTTAGTCACCGTCTCCTCA	27
(2) INFORMATION FOR SEQ ID NO:10:	
(i) SEQUENCE CHARACTERISTICS:  (A) LEMGTH: 27 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:	
AATAGATATCTCCTTCAACACCTGCAA	27
(2) INFORMATION FOR SEQ ID NO:11:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:	
ATCGGGACAAAGTTGGAAATA	21
(2) INFORMATION FOR SEQ ID NO:12:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:	

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GGCGGTCTGGTACCGG

(2) INFORMATION FOR SEQ ID NO:13:

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDMESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:	
GTCAACAACATAGTCATCA	19
(2) INFORMATION FOR SEQ ID NO:14:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDMESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:	
CACAGGTGTGTCCCCAAGGAAAA	23
(2) INFORMATION FOR SEQ ID NO:15:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 18 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:	
AATCTGGGGTAGGCACAA	18
(2) INFORMATION FOR SEQ ID NO:16:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDMESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:	
AGTGTGTGTCCCCAAGG	17



(2) INFORMATION FOR SEQ ID NO:17:

(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear		
(ii)	MOLECULE TYPE: cDNA		
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:	17:	
CACAGCTGCC	CGCCCAGGTGGCAT	24	
(2) INFORM	ATION FOR SEQ ID NO:18:		
(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDMESS: single (D) TOPOLOGY: linear		
(ii)	MOLECULE TYPE: cDNA		
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:	18:	
GTCGCCAGTG	CTCCCTT	17	
(2) INFORM	ATION FOR SEQ ID NO:19:		
(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear		
(ii)	MOLECULE TYPE: cDNA		
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:	19:	
ATCGGACGTG	GACGTGCAGA	20	
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